Applicant: Clancy et al.

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of claims:

- (presently amended) A method of diagnosing or determining the risk of developing gastric cancer in a subject with a *Helicobacter* infection, including:
 - a) determination of IgG2 anti-H. pylori antibody level in the subject; and
 - b) comparison of the IgG2 anti-*H. pylori* antibody level with a predetermined control IgG2 anti-*H. pylori* antibody level, wherein a reduction in the level of IgG2 anti-*H. pylori* antibody in the subject compared to the control indicates the presence of gastric cancer or an increased risk of developing gastric cancer.
- 2-22. (presently cancelled)
- 23. (added) A method of diagnosing or determining the risk of developing gastric cancer in a subject with a *Helicobacter* infection, including:
 - a) determination of yIFN level in the subject;
 - b) comparison of the γIFN level with a predetermined control γIFN level, wherein a reduction in the level of γIFN in the subject compared to the control indicates the presence of gastric cancer or an increased risk of developing gastric cancer.
- 24. (added) A method of diagnosing or determining the risk of developing gastric cancer in a subject with a *Helicobacter* infection, including:
 - a) determination of IL-4 level in the subject;
 - b) comparison of the IL-4 level with a predetermined control IL-4 level, wherein an elevation in the level of IL-4 in the subject compared to the control indicates the presence of gastric cancer or an increased risk of developing gastric cancer.
- 25. (added) A method of diagnosing or determining the risk of developing gastric cancer in a subject with a *Helicobacter* infection, including a combination of a method according to claim 1 and a method according to claim 23.
- 26. (added) A method of diagnosing or determining the risk of developing gastric cancer in a subject with a *Helicobacter* infection, including a combination of a method according to claim 1 and a method according to claim 24.
- 27. (added) The method of claim 25, further comprising a method according to claim 24.

Applicant: Clancy et al.

- 28. (added) A method of diagnosing or determining the risk of developing gastric cancer in a subject with a *Helicobacter* infection, including a combination of a method according to claim 23 and a method according to claim 24.
- 29. (added) A method according to any one of claims 1, 23 or 24 wherein the *Helicobacter* infection is a *Helicobacter pylori* infection.
- 30. (added) A method according to claim 1 wherein the IgG2 anti-*H. pylori* antibody level is determined by detection of the level in a sample of biological fluid.
- 31. (added) A method according to claim 23 wherein the γIFN level is determined by detection of the level in a sample of biological fluid.
- 32. (added) A method according to claim 24 wherein the IL-4 level is determined by detection of the level in a sample of biological fluid.
- 33. (added) A method according to claim 1 wherein the IgG2 anti-*H. pylori* antibody level is determined by detection of the level in a biological fluid selected from the group consisting of blood, saliva and gastric fluid.
- 34. (added) A method according to claim 23 wherein the γIFN level is determined by detection of the level in a biological fluid selected from the group consisting of blood, saliva and gastric fluid.
- 35. (added) A method according to claim 24 wherein the IL-4 level is determined by detection of the level in a biological fluid selected from the group consisting of blood, saliva and gastric fluid.
- 36. (added) A method according to claim 1 wherein the determination of the IgG2 anti-H. pylori antibody level either simultaneously provides, or can be performed simultaneously with, a method which provides an indication of H. pylori status.
- 37. (added) A method according to claim 23 wherein the determination of the γIFN level either simultaneously provides, or can be performed simultaneously with, a method which provides an indication of *H. pylori* status.
- 38. (added) A method according to claim 24 wherein the determination of the IL-4 level either simultaneously provides, or can be performed simultaneously with, a method which provides an indication of *H. pylori* status.

Attorney Docket No. 24356-002 (BSW-2)

Applicant: Clancy et al.

- 39. (added) A method according to claim 1 wherein the IgG2 anti-H. pylori antibody level is detected by a near-subject assay.
- (added) A method according to claim 23 wherein the γIFN level is detected by a near-subject assay.
- 41. (added) A method according to claim 24 wherein the IL-4 level is detected by a near-subject assay.
- 42. (added) A method according to claim 1 wherein the IgG2 anti-*H. pylori* antibody level is determined by an antibody assay.
- 43. (added) A method according to claim 23 wherein the γIFN level is determined by an antibody assay.
- 44. (added) A method according to claim 24 wherein the IL-4 level is determined by an antibody assay.
- 45. (added) A method according to claim 1 wherein the IgG2 anti-H. pylori antibody level is determined by ELISA.
- 46. (added) A method according to claim 23 wherein the γIFN level is determined by ELISA.
- 47. (added) A method according to claim 24 wherein the IL-4 level is determined by ELISA.
- 48. (added) A method of predicting the risk of, or diagnosing, gastric cancer in a subject having a *Helicobacter* infection by
 - a) determining the frequency of IgG2 anti-H. pylori antibody- and/or γIFN- and/or IL-4-producing cells in the subject's blood; and
 - b) comparison of the frequency of IgG2 anti-*H. pylori* antibody- and/or γIFN- and/or IL-4-producing cells in the subject's blood with a predetermined control level, wherein a reduction in the level of IgG2 anti-*H. pylori* antibody- and/or γIFN-producing cells and/or an elevation in IL-4-producing cells in the subject's blood indicates the presence of gastric cancer or an increased risk of developing gastric cancer.
- 49. (added) A method according to claim 48 wherein the blood is purified to provide an enriched white blood cell population.
- 50. (added) A method according to claim 48 wherein the blood is purified to provide an

Applicant: Clancy et al.

enriched white blood cell population and the white blood cell population is further fractionated to obtain specific cell populations.

- 51. (added) A method according to claim 48 wherein when the frequency of IgG2 anti-H. pylori antibody-producing cells is determined, the IgG2 anti-H. pylori antibody-producing cells are stimulated with H. pylori antigen prior to determination of the frequency of IgG2 anti-H. pylori antibody-producing cells.
- 52. (added) A method according to claim 48 wherein when the frequency of γIFN-producing cells is determined, the γIFN-producing cells are stimulated with *H. pylori* antigen prior to determination of the frequency of γIFN-producing cells.
- 53. (added) A method according to claim 48 wherein when the frequency of IL-4-producing cells is determined, the IL-4-producing cells are stimulated with *H. pylori* antigen prior to determination of the frequency of IL-4-producing cells.
- 54. (added) A method of predicting the risk of, or diagnosing, gastric cancer in a subject having a *Helicobacter* infection by
 - a) determining the frequency of IgG2 anti-H. pylori antibody and/or γIFN and/or
 IL-4-producing cells in the subject's gastric mucosa; and
 - b) comparison of the frequency of IgG2 anti-*H. pylori* antibody and/or γIFN and/or IL-4-producing cells in the subject's gastric mucosa with a predetermined control level, wherein a reduction in the level of IgG2 anti-*H. pylori* antibody-and/or γIFN-producing cells and/or an elevation in IL-4-producing cells in the subject's gastric mucosa indicates the presence of gastric cancer or an increased risk of developing gastric cancer.
- 55. (added) A method according to claim 54 wherein the cells are derived from a biopsy sample.
- 56. (added) A method according to claim 54 wherein the frequency of IgG2 anti-*H. pylori* antibody- and/or γIFN- and/or IL-4-producing cells is determined by flow cytometry.